- For receiving Office use only International Application No. REQUEST International Filing Date The undersigned requests that the present international application be processed Name of receiving Office and "PCT International Application" according to the Patent Cooperation Treaty. (if desired) (12 characters maximum) NEB-236-PCT TITLE OF INVENTION Box No. I MODULATION OF ENZYME ACTIVITY BY CHANGING RECIPROCAL STEREO-GEOMETRIC POSITIONS OF TWO CATALYTIC CENTERS IN AN ENZYME Box No. II APPLICANT This person is also inventor Name and address: (Family name followed by given name; for a legal entity, full official designation. The address must include postal code and name of country. The country of the address indicated in this Box is the applicant's State (that is, country) of residence if no State of residence is indicated below.) Telephone No. Facsimile No. NEW ENGLAND BIOLABS, INC. 32 Tozer Road Teleprinter No. Beverly, MA 01915 US Applicant's registration No. with the Office State (that is, country) of residence: State (that is, country) of nationality: US the United States of America only the States indicated in the Supplemental Box This person is applicant for the purposes of: all designated States all designated States except the United States of America FURTHER APPLICANT(S) AND/OR (FURTHER) INVENTOR(S) Box No. III Name and address: (Family name followed by given name; for a legal entity, full official designation The address must include postal code and name of country. The country of the address indicated in this This person is: Box is the applicant's State (that is, country) of residence if no State of residence is indicated below.) applicant only GUAN, Chudi applicant and inventor 10 Great Pond Road inventor only (If this check-box Wenham, MA 01984 is marked, do not fill in below.) US Applicant's registration No. with the Office State (that is, country) of residence: State (that is, country) of nationality: the States indicated in the Supplemental Box This person is applicant for the purposes of: all designated States except the United States of America the United States all designated of America only Further applicants and/or (further) inventors are indicated on a continuation sheet. AGENT OR COMMON REPRESENTATIVE; OR ADDRESS FOR CORRESPONDENCE Box No. IV The person identified below is hereby/has been appointed to act on behalf of the applicant(s) before the competent International Authorities as: common representative agent Telephone No. Name and address: (Family name followed by given name; for a legal entity, full official designation. The address must include postal code and name of country.) 978-927-5054 STRIMPEL, Harriet M. Facsimile No. New England Biolabs, Inc. 32 Tozer Road Teleprinter No. Beverly, MA 01915 US Agent's registration No. with the Office

Address for correspondence: Mark this check-box where no agent or common representative is/has been appointed and the space above is used instead to indicate a special address to which correspondence should be sent.

Addendum

1. Modulation of Enzyme Activity by Changing Reciprocal Stereo-Geometric Positions of Two Catalytic Centers in an Enzyme

Continuation of Box No. III FURTHER APPLICANT(S) AND/OR (FURTHER). INVENTOR(S)					
If none of the following sub-boxes is a	used, this sheet should not be	included in the requ	iest.		
Name and address: (Family name followed by given name; for a legal entity, full official designation. The address must include postal code and name of country. The country of the address indicated in this Box is the applicant's State (that is, country) of residence if no State of residence is indicated below.) KUMAR, Sanjay 83 Old Right Road Ipswich, MA 01938 US			This person is: applicant only X applicant and inventor inventor only (If this check-box is marked, do not fill in below.) Applicant's registration No. with the Office		
State (that is, country) of nationality:		State (that is, country)	of residence:		
US .		JS			
This person is applicant for the purposes of:	signated all designated S s the United State	tates except s of America	the United States the States indicated in the Supplemental Box		
Name and address: (Family name followed The address must include postal code and name to be a six is the applicant's State (that is, country) of KUCERA, Rebecca 160 Woodland Mead Hamilton, MA 01982 US	me of country. The country of the a	ddress indicated in this	This person is: applicant only X applicant and inventor inventor only (If this check-box is marked, do not fill in below.) Applicant's registration No. with the Office		
State (that is, country) of nationality:		State (that is, country	y) of residence:		
This person is applicant for the purposes of:	esignated all designated	US States except X	the United States the States indicated in the Supplemental Box		
Name and address: (Family name followe. The address must include postal code and na Box is the applicant's State (that is, country) o	ame of country. The country of the	address indicated in this	This person is: applicant only applicant and inventor inventor only (If this check-box is marked, do not fill in below.) Applicant's registration No. with the Office		
State (that is, country) of nationality:		State (that is, countr	y) of residence:		
This person is applicant all of or the purposes of:	designated all designated the United Sta	States except tes of America	the United States of America only the States indicated in the Supplemental Box		
Name and address: (Family name follows The address must include postal code and n Box is the applicant's State (that is, country)	ame of country. The country of the	address indicated in this			
State (that is, country) of nationality	<i>r</i> .	State (that is, count	ry) of residence:		
		d States except ates of America	the United States of America only the Supplemental Box		
Further applicants and/or (further) inventors are indicated on another continuation sheet.					

Sheet No. ...3...

Box No. V DESIGNATIO	ONS					
The filing of this request constitutes under Rule 4.9(a), the designation of all Contracting States bound by the PCT on the international filing date, for the grant of every kind of protection available and, where applicable, for the grant of both regional and national patents.						
However,						
DE Germany is not design	gnated for any kind of nationa	al protection				
KR Republic of Korea is	not designated for any kind	of national protection				
RU Russian Federation i	s not designated for any kind	of national protection				
(The check-boxes above may be used to exclude (irrevocably) the designations concerned in order to avoid the ceasing of the effect, under the national law, of an earlier national application from which priority is claimed. See the Notes to Box No. V as to the consequences of such national law provisions in these and certain other States)						
Box No. VI PRIORITY	CLAIM					
The priority of the following ea	arlier application(s) is hereby	claimed:				
Filing date	Number	Where earlier application is:				
of earlier application (day/month/year)	of earlier application	national application: country or Member of WTO	regional application:* regional Office	international application: receiving Office		
item (1) 21 November 2003 (21.11.2003)	60/524,123	US				
item (2)						
item (3)						
				<u> </u>		
Further priority claims a	re indicated in the Supplemer	ntal Box.				
The receiving Office is requested to prepare and transmit to the International Bureau a certified copy of the earlier application(s) (only if the earlier application was filed with the Office which for the purposes of this international application is the receiving Office) identified above as:						
all items X item (1) item (2) item (3) other, see Supplemental Box * Where the earlier application is an ARIPO application, indicate at least one country party to the Paris Convention for the Protection of Industrial Property or one Member of the World Trade Organization for which that earlier application was filed (Rule 4.10(b)(ii)):						
Industrial Property of One M	oj ine moriu Truue Or					
Box No. VII INTERNA	ATIONAL SEARCHING AU	THORITY				
Choice of International Searching Authority (ISA) (if two or more International Searching Authorities are competent to carry out the international search, indicate the Authority chosen; the two-letter code may be used):						
ISA /US						
Request to use results of earlier search; reference to that search (if an earlier search has been carried out by or requested from the International Searching Authority):						
Date (day/month/year)	Num	ber Cou	ıntry (or regional Office))		
Box No. VIII DECLA	Box No. VIII DECLARATIONS					
	are contained in Boxes Nos. cate in the right column the n			Number of declarations		
Box No. VIII (i)	Box No. VIII (i) Declaration as to the identity of the inventor :					
Box No. VIII (ii) Declaration as to the applicant's entitlement, as at the international filing date, to apply for and be granted a patent:						
Box No. VIII (iii)	Box No. VIII (iii) Declaration as to the applicant's entitlement, as at the international filing date, to claim the priority of the earlier application :					
Box No. VIII (iv) Declaration of inventorship (only for the purposes of the designation of the United States of America)						
Box No. VIII (v)	Declaration as to non-prej	udicial disclosures or exc	eptions to lack of novelty	y :		

Sheet	No	4

Box No. IX CHECK LIST; LANGUAGE C	of FILING				
This international application contains: (a) in paper form, the following number of sheets:	This international application is accompanied by the following item(s) (mark the applicable check-boxes below and indicate in right column the number of each item):	Number of items			
request (including	1. fee calculation sheet	:			
declaration sheets) : 4	2. original separate power of attorney	:			
description (excluding sequence listings and/or	3. original general power of attorney	:			
tables related thereto) : 38	4. copy of general power of attorney; reference number,				
claims : 6	if any:	:			
abstract : 1	5. statement explaining lack of signature	•			
drawings : 22 Sub-total number of sheets : 71	6. priority document(s) identified in Box No. VI as item(s):	:			
sequence listings :	7. translation of international application into (language):	:			
tables related thereto : (for both, actual number of	8. separate indications concerning deposited microorganism or other biological material	:			
sheets if filed in paper form, whether or not also filed in computer readable form;	9. X sequence listings in computer readable form (indicate type and number of carriers)				
see (c) below)	(i) Copy submitted for the purposes of international search under Rule 13ter only (and not as part of the international application)	n):12			
Total number of sheets : 71 (b) only in computer readable form (Section 801(a)(i))	(ii) (only where check-box (b)(i) or (c)(i) is marked in left column additional copies including, where applicable, the copy for the purposes of international search under Rule 13ter	<i>y</i> •			
(i) sequence listings	(iii) X together with relevant statement as to the identity of the copy copies with the sequence listings mentioned in left column	or : 2			
(ii) tables related thereto (c) also in computer readable form	10. tables in computer readable form related to sequence listings (indicate type and number of carriers)				
(Section 801(a)(ii)) (i) sequence listings	(i) copy submitted for the purposes of international search under Section 802(b-quater) only (and not as part of the internation	al			
(ii) tables related thereto	application)	:			
Type and number of carriers (diskette, CD-ROM, CD-R or other) on which are contained the	(ii) (only where check-box (b)(ii) or (c)(ii) is marked in left column additional copies including, where applicable, the copy for the purposes of international search under Section 802(b-quater)	ne) :			
sequence listings:	(iii) together with relevant statement as to the identity of the copy	or .			
tables related thereto:	copies with the tables mentioned in left column	:1			
(additional copies to be indicated under items 9(ii) and/or 10(ii), in right column)	11. \(other (specify):\$2,946.check; return post-card	• •			
	Caru				
Figure of the drawings which should accompany the abstract: 12	Language of filing of the international application: English				
Box No. X SIGNATURE OF APPLICA	NT, AGENT OR COMMON REPRESENTATIVE	the request)			
Next to each signature, indicate the name of the person signature.	ening and the capacity in which the person signs (if such capacity is not obvious from reading	me requesty.			
	# Zmest O hungs of	-			
	Harriet M. Strimpel, Chief Patent Counse	l and			
	Assistant Clerk				
	For receiving Office use only				
Date of actual receipt of the purported international application:	Date of actual receipt of the purported international application: 2. Drawings:				
3. Corrected date of actual receipt due to later but timely received papers or drawings completing the purported international application:					
4. Date of timely receipt of the required corrections under PCT Article 11(2):		not received:			
5. International Searching Authority (if two or more are competent): ISA /	6. Transmittal of search copy delayed until search fee is paid				
For International Bureau use only					
Date of receipt of the record copy					
by the International Bureau:					

This sheet is not part of and does not count as a sheet of the international application.

PCT For receiving Office use only FEE CALCULATION SHEET International Application No. Annex to the Request Applicant's or agent's NEB-236-PCT file reference Date stamp of the receiving Office Applicant NEW ENGLAND BIOLABS, INC., et al. CALCULATION OF PRESCRIBED FEES 300.00 T 1. TRANSMITTAL FEE..... 1,000.00 S 2. SEARCH FEE.... International search to be carried out by (If two or more International Searching Authorities are competent to carry out the international search, indicate the name of the Authority which is chosen to carry out the international search.) 3. INTERNATIONAL FILING FEE Where item (b) and/or (c) of Box No. IX apply, enter Sub-total number of sheets Where item (b) and (c) of Box No. IX do not apply, enter Total number of sheets 1,134.00 ii 492.00 number of sheets additional component (only if sequence listings and/or tables related thereto are filed in computer readable form under Section 801(a)(i), or both in that form and on paper, under Section 801(a)(ii)): i3 1,626.00 Add amounts entered at i1, i2 and i3 and enter total at I (Applicants from certain States are entitled to a reduction of 75% of the international filing fee. Where the applicant is (or all applicants are) so entitled, the total to be entered at I is 25% of the international filing fee.) 20.00 P 4. FEE FOR PRIORITY DOCUMENT (if applicable)..... **USD** 2,946.00 Add amounts entered at T, S, I and P, and enter total in the TOTAL box MODE OF PAYMENT authorization to charge deposit account (see below) postal money order cash coupons X cheque bank draft revenue stamps ___ other (specify): AUTHORIZATION TO CHARGE (OR CREDIT) DEPOSIT ACCOUNT Receiving Office: RO/ (This mode of payment may not be available at all receiving Offices) Deposit Account No.: 14-0740 Authorization to charge the total fees indicated above. Date: 22 November 2004 (This check-box may be marked only if the conditions for deposit accounts of the receiving Office so permit) Authorization to charge any deficiency Name: Harriet M. Strimpel or credit any overpayment in the total fees indicated above. Authorization to charge the fee for priority document. Signature: